

Docket No. 20488/57

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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

APPLICANT:

Kühn-Wache et al.

ART UNIT:

Not yet assigned

SERIAL NO.:

10/667,200

FILED:

September 18, 2003

FOR: SECONDARY BINDING SITE OF DIPEPTIDYL PEPTIDASE

IV (DP IV)

CERTIFICATE OF MAILING

I hereby certify that this paper (along with any paper referred to as being attached or enclosed) is being deposited with the United States Postal Service on the date shown below with sufficient postage as first class mail in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

By: Sandra Wittup
Sandra J. Wittrup

November 19,2003

Date

INFORMATION DISCLOSURE STATEMENT SUBMITTING REFERENCES ON COMPACT DISC

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

Date: November 19, 2003

Pursuant to Applicant(s) duty of disclosure, the information listed in the attached form PTO-1449 is brought to the attention of the Examiner.

The citation of the listed items is not a representation that they constitute a complete or exhaustive listing of the relevant art or that the references are prior art. The items listed are submitted in good faith, but are not intended to substitute for the Examiner's search. It is hoped, however, that in addition to apprising the Examiner of these particular items, they will assist in identifying fields of search and in making as full and complete a search as possible.

The filing of this information disclosure statement is not an admission that the information cited herein is, or is considered to be, material to patentability as defined in 37 C.F.R. § 1.56(b).

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In accordance with the requirement under 37 C.F.R. 1.98 (3)(i), the following are concise explanations of the relevance, as presently understood, with regard to those items submitted herewith that are not in the English language:

I. LIST OF PATENTS, PUBLICATIONS OR OTHER INFORMATION

The patents, publications or other information submitted for consideration by the Office are listed on PTO-1449, attached hereto.

II. COPIES

a. 🔯 Submitted herewith on Compact Disc is a legible copy of (i) each U.S. and foreign patent; (ii) each publication or that portion which caused it to be listed; and (iii) all other information or that portion which caused it to be listed.

III. CONCISE EXPLANATION OF THE RELEVANCE

(check at least one box)

- Except as may be indicated below in (b), all of the patents, publications or a. 🔲 other information are in the English language or were cited in an English language Search Report, a copy of which is attached hereto (concise explanation not required).
- b. 🖂 A concise explanation of the relevance of all patents, publications or other information listed that is not in the English language is as follows:

FRENCH LANGUAGE FR 2 696 740 Applicants have relied on an English language abstract in determining that this patent apparently relates to Dimethylbiguanide drug derivatives and their medical applications.

FRENCH LANGUAGE FR 2 085 665 Applicants have relied on an English language abstract in determining that this patent apparently relates to biguanide substitutes possessing a hypoglycemic property.

GERMAN LANGUAGE DT 25 42 598 A1 Applicants have relied on an English language abstract in determining that this patent apparently relates to biguanide salts as well as a method for preparing such salts. In addition to this the invention apparently covers pharmaceutical compounds which contain such salts.

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GERMAN LANGUAGE WO 97/40832 Applicants have relied on an English language abstract in determining that this patent apparently relates to the use of a method for reducing in the blood of a mammal by administration of effectors the enzyme activity of dipeptidyl peptidase (DP IV) or enzyme activity similar to DP IV, the endogenous (or additionally exogenously administered) insulinotropic peptide gastric inhibitory polypeptide 1-42 and glucagon-like peptide amide-1 7-36 are decomposed in a causal sequence to a reduced extent by DP IV enzymes or those similar to DP IV.

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JAPANESE PATENT JP 4-334357A2: Applicants have relied on an English language abstract in determining that this patent ACYL DERIVATIVE HAVING ENZYME-INHIBITING ACTION, apparently relates to a compound having a prolyl endopeptibase activity-inhibiting action and useful as an antidement agent, especially an anti-amnestic agent.

GERMAN LANGUAGE DE 299 09 210 U1 Applicants have relied on an English language abstract in determining that this patent apparently relates to dipeptide compounds or compounds analogous to dipeptide compounds, which are made of an amino acid and a thiazolidine or pyrrolidine group, and to their salts. The invention further relates to the use of these compounds in the treatment of impaired glucose tolerance, glucosuria, hyperlipidemia, metabolic acidosis, diabetes mellitus, diabetic neuropathy and nephropathy as well as secondary diseases of diabetes mellitus in mammals.

GERMAN LANGUAGE DE 198 26 972 A1 Applicants have relied on an English language abstract in determining that this patent apparently relates to the use of inhibitors (I) of dipeptidyl peptidase IV (DP IV) or analogous enzymes for inhibiting the activation and proliferation of keratinocytes (including malignantly transformed keratinocytes) and tumor cells derived from keratinocytes.

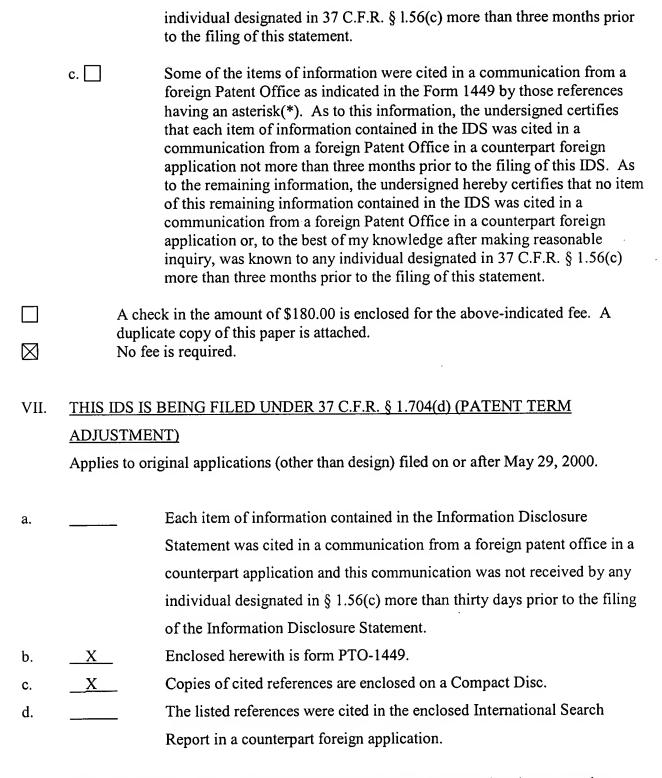
GERMAN LANGUAGE WO 99/61431 Applicants have relied on an English language abstract in determining that this patent apparently relates to dipeptide compounds or compounds analogous to dipeptide compounds, which are made of an amino acid and a thiazolidine or pyrrolidine group, and to their salts. The invention further relates to the use of these compounds in the treatment of impaired glucose tolerance, glucosuria, hyperlipidemia, metabolic acidosis, diabetes mellitus, diabetic neuropathy and nephropathy as well as secondary diseases of diabetes mellitus in mammals.

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| | c. 🗌 | The following additional information is provided for the Examiner's consideration: | | | | | |
|-------|-----------------------------------|---|--|--|--|--|--|
| | | <u>FEES</u> | | | | | |
| IV. | THIS IDS IS (check one bo | BEING FILED UNDER 37 C.F.R. § 1.97(b) | | | | | |
| | a. 🔀 | within three months of the filing date of a national application (37 C.F.R. § 1.97(b) (1)). No fee or certification is required. | | | | | |
| | b. 🗌 | within three months of the date of entry of the national stage as set forth in §1.491 in an international application (37 C.F.R. § 1.97(b) (2)). No fee or certification is required. | | | | | |
| | с. 🗌 | before the mailing date of a first Action on the merits (37 C.F.R. § 1.97(b) (3)). No fee or certification is required. In the event that a first Office Action on the merits has been issued, please consider this IDS under 37 C.F.R. § 1.97(c) and see the certification under 37 C.F.R. § 1.97(e)below, or, if no certification has been made, charge our deposit account a fee in the amount of \$180.00 as required by 37 C.F.R. § 1.17(p). | | | | | |
| V. | THIS IDS IS (check one bo | BEING FILED UNDER 37 C.F.R. § 1.97(c): (bx) | | | | | |
| | 1.97(c) (1)) o | tiling date of a Final Office Action under 37 C.F.R. § 1.113 (See 37 C.F.R. § r before the mailing date of a Notice of Allowance under 37 C.F.R. § 1.311 R. § 1.97(c) (2)). | | | | | |
| | a. No ce | rtification; therefore, a fee in the amount of \$180.00 is required by 37 C.F.R. § 1.17(p). | | | | | |
| | b. See th | or the certification below. No fee is required. | | | | | |
| VI. | CERTIFICAT | TION UNDER 37 C.F.R. § 1.97(e) (check only one box) | | | | | |
| The u | undersigned hereby certifies that | | | | | | |
| | a. 🗌 | each item of information contained in the IDS was cited in a communication from a foreign Patent Office in a counterpart foreign application not more than three months prior to the filing of this IDS; or | | | | | |
| | b. 🔲 | no item of information contained in the IDS was cited in a communication from a foreign Patent Office in a counterpart foreign application or, to the best of my knowledge after making reasonable inquiry, was known to any | | | | | |

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If the Examiner has any questions concerning this IDS, the Examiner is requested to contact the undersigned. If it is determined that this IDS has been filed under the wrong rule, the

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PTO is requested to consider this IDS under the proper rule (with a petition, if necessary) and charge any additional fees to Deposit Account No. 50-0369.

Date: November 19, 2003

Respectfully submitted,

John C. Serio, Reg. No. 39,023

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| INFORMATI | ON DISCLOSURE STATEMENT |

ATTY. DOCKET NO. **20488/57**

SERIAL NO. 10/667,200

APPLICANT(S): Kühn-Wache et al.

FILING DATE: September 18, 2003 ART UNIT:
Not Yet Assigned

UNITED STATES PATENT DOCUMENTS

• . • . •

| EXAM. INITIAL | | DOCUMENT NUMBER | DATE | INVENTOR | CLASS | SUB CLASS | FIL. DATE IF APPR |
|------------------|----|--------------------|------------|-------------------|-------|--------------|-------------------------|
| | AA | 2,961,377 | 11/22/1960 | Shapiro et al. | 167 | 65 | |
| | AB | 3,174,901 | 03/23/1965 | Sterne | 167 | 65 | |
| | AC | 3,879,541 | 04/22/1975 | Kabbe et al. | 424 | 326 | |
| | AD | 3,960,949 | 06/01/1976 | Ahrens et al. | 260 | 564 B | |
| | AE | 4,028,402 | 06/07/1977 | Fischer et al. | 260 | 501.14 | |
| | AF | 4,935,493 | 06/19/1990 | Bachovchin et al. | 530 | 331 | |
| | AG | 5,433,955 | 07/18/1995 | Bredehorst et al. | 424 | 94.3 | |
| | AH | 5,462,928 | 10/31/1995 | Bachovchin et al. | 514 | 19 | |
| | ΑI | 5,512,549 | 04/30/1996 | Chen et al. | 514 | 12 | |
| | AJ | 5,543,396 | 08/06/1996 | Powers et al. | 514 | 19 | |
| | AK | 5,614,379 | 03/25/1997 | MacKellar | 435 | 68.1 | |
| | AL | 5,624,894 | 04/29/1997 | Bodor | 514 | 2 | |
| | AM | 5,939,560 | 08/17/1999 | Jenkins et al. | 548 | 535 | |
| | AN | 6,006,753 | 12/28/1999 | Efendic | 128 | 898 | |
| | AO | 5,827,898 | 10/27/1998 | Khandwala et al. | 514 | 734 | |
| | AP | 6,201,132 B1 | 03/13/2001 | Jenkins et al. | 548 | 535 | 04/21/99 |
| | AQ | 6,303,661 B1 | 10/16/2001 | Demuth et al. | 514 | 866 | 04/24/97 |
| | AR | 6,500,804 B2 | 12/31/2002 | Demuth et al. | 514 | 19 | 12/31/02 |
| | AS | 6,319,893 B1 | 11/20/2001 | Demuth et al. | 514 | 2 | 11/20/01 |
| | AT | 6,011,155 | 01/04/2000 | Villhauer | 544 | 333 | |
| | AU | 6,107,317 | 08/22/2000 | Villhauer | 514 | 365 | |
| | AV | 6,110,949 | 08/29/2000 | Villhauer | 514 | 365 | |
| | AW | 6,124,305 | 09/26/2000 | Villhauer | 514 | 272 | |
| | AX | 6,172,081 | 01/09/2001 | Damon | 514 | 307 | |
| Examiner: | | | | Date: | | | |

| | | ATEMENT | 20488/57 | | 10/667,200 | |
|-----|--------------------|--------------|------------------------------------|-------|--------------------------------|------------------|
| | | • | APPLICANT(S): Kühn-Wache et al. | | | |
| | | | FILING DATE: September 18, 2003 | | ART UNIT: Yet Assign | |
| | | UNITED STATE | ES PATENT DOCUMENTS | | | , |
| AY | 5,705,483 | 01/06/1998 | Galloway et al. | 514 | 12 | |
| AZ | 2001/0025023 A1 | 09/27/2001 | Carr | 514 | 2 | |
| AAA | 6,548,481 B1 | 04/15/2003 | Demuth et al. | 514 | 19 | 11/28/00 |
| | | FOREIGN I | PATENT DOCUMENTS | | | |
| | DOCUMENT NUMBER | DATE | COUNTRY | CLASS | SUB CLASS | TRAN Y/N |
| BA | WO 01/62266 A2 | 08/30/2001 | PCT | A61K | 38/00 | Y |
| ВВ | WO 00/53171 | 09/14/2000 | PCT | A61K | 31/155 | Y |
| BC | DT 25 42 598 A1 | 04/22/1976 | Germany | C07C | 129/16 | N |
| BD | FR 2 696 740 A1 | 04/15/1994 | France | C07D | 207/404 | N |
| BE | FR 2 085 665 | 12/31/1971 | France | A61K | 27/00 | N |
| BF | WO 97/40832 | 11/06/1997 | PCT | A61K | 31/425 | Abstract only |
| BG | WO 95/15309 | 06/08/1995 | PCT | C07D | 207/16 | Y |
| ВН | JP 4334357 | 11/20/1992 | Japan | C07C | 233/57 | Abstract only |
| BI | WO 93/08259 | 04/29/1993 | PCT | C12N | | Y |
| ВЈ | WO 95/11689 | 05/04/1995 | PCT | A61K | 37/00 | Y |
| BK | WO 97/45117 | 12/04/1997 | PCT | A61K | 31/435 | Y |
| BL | DE 196 16 486 C2 | 10/30/1997 | Germany | A61K | 45/00 | Y |
| ВМ | WO 95/29691 | 11/09/1995 | PCT | A61K | 38/00 | Y |
| BN | WO 98/22494 | 05/28/1998 | PCT | C07K | 5/06 | Y |
| ВО | WO 00/01849 | 01/13/2000 | PCT | C12Q | 1/68 | Y |
| BP | EP 0 658 568 A1 | 06/21/1995 | EPO | C07K | 14/605 | Y |
| BQ | DD 296 075 A5 | 11/21/1991 | Germany | C07D | 295/04 | Y |
| BR | DD 296 075 A5 | 11/21/1991 | Germany | C07D | 295/04 | N |
| BS | EP 0 708 179 A2 | 04/24/1996 | EPO | C12N | 15/16 | Y |

| INFOR | FORM PTO-1449 MATION DISCLOSURE ST | TATEMENT | ATTY. DOCKET NO. SERIAL NO. 10/667,200 | | | |
|----------|---------------------------------------|------------|--|-------|-------------------------|--------------------|
| | | | APPLICANT(S): Kühn-Wache et al. | | | |
| | | | FILING DATE: September 18, 2003 | 1 | ART UNIT: Yet Assign | ed |
| | | FOREIGN I | PATENT DOCUMENTS | | | |
| | DOCUMENT NUMBER | DATE | COUNTRY | CLASS | SUB CLASS | TRAN Y/N |
| B | Γ EP 0 995 440 A1 | 04/26/2000 | EPO | A61K | 31/425 | N |
| В | J WO 91/11457 | 08/08/1991 | PCT | C07K | 7/34 | Y |
| В | WO 91/16339 | 10/31/1991 | PCT | C07K | 5/10 | Y |
| В | W WO 98/19998 | 05/14/1998 | PCT | C07D | 207/00 | Y |
| B | WO 91/17767 | 11/28/1991 | PCT | A61K | 37/54 | Y |
| В | Y JP 04-288098 | 10/13/1992 | JР | A61K | 37/64 | (Abstract Only) |
| В | Z WO 99/46272 A | 09/16/1999 | PCT | C07F | 9/572 | Y |
| ВА | A DE 299 09 210 U | 09/09/1999 | Germany | A61K | 31/425 | N |
| BA | B DE 198 26 972 A1 | 12/23/1999 | Germany | A61K | 38/05 | N |
| BA | C WO 01/97808 | 12/27/2001 | PCT | A61K | 31/425 | Y |
| ВА | D WO 01/34594 A1 | 05/17/2001 | PCT | C07D | 401/06 | Y |
| BA | E WO 00/10549 | 03/02/2000 | PCT | A61K | 31/00 | Y |
| BA | AF WO 01/74299 A2 | 10/11/2001 | PCT | A61K | | Y |
| BA | G WO 02/20825 A1 | 03/14/2002 | PCT . | C12Q | 1/00 | Y |
| BA | H WO 01/89569 A1 | 11/29/2001 | PCT | A61K | 45/06 | Y |
| BA | AI WO 99/62914 | 12/09/1999 | PCT | C07F | 5/02 | Y |
| BA | AJ WO 99/61431 | 12/02/1999 | PCT | C07D | 277/04 | Abstract Only- |
| BA | K EP 1 130 022 A1 | 09/15/2001 | ЕРО | C07D | 498/09 | Y |
| BA | AL WO 01/32624 A1 | 05/10/2001 | PCT | C07D | 217/04 | Y |
| BA | M WO 01/09169 A2 | 02/08/2001 | PCT | C07K | 5/062 | Y |
| BA | N WO 01/94310 A1 | 12/13/2001 | PCT | C07D | 209/12 | Y |
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| INFORM | FORM PTO-1449 ATION DISCLOSURE STATEMENT | ATTY. DOCKET NO. 20488/57 | SERIAL NO. 10/667,200 | |
|--------|---|---|---|--|
| | | APPLICANT(S): Kühn-Wache et al. | | |
| | | FILING DATE: September 18, 2003 | ART UNIT: Not Yet Assigned | |
| | OTHER DOCUMENTS (INCLUDING A | AUTHOR, TITLE, DATE, PERTINEN | IT PAGES, ETC.) | |
| CA | Campbell, I.W. New Antidiabetic Drugs, e efficacy and inadequacy". 3:33-51 (1990). | d. C.J. Bailey & P.R. Flatt, Smith-Gordo | n, "Sulphonylureas and metformi | |
| СВ | CHEMICAL ABSTRACTS, vol. 115. No. 149947q, SCHOEN EKKEHARD ET AL: inhibitors on activity of dipeptidyl peptidas | "Dipeptidyl peptidase IV in the immune | system. Effects of specific enzyr | |
| CC | CHEMICAL ABSTRACTS, vol. 118, no. 2 Hosoda, et al, "Preparation of N-(heterocyc November 1992 (1992-11-20) | 25, 21. June 1993 (1993-06-21) Columbu lic Carbonyl) Amino Acids and Analogs | s, Ohio, US; abstract no. 2553421 as Prolyl Endopeptidase Inhibito | |
| CD | CHEMICAL ABSTRACTS, vol. 126, no. 2, 13. January 1997 (1997-01-13) Columbus, Ohio, US; abstract no. 161 STOECKEL A. ET AL: "Competitive inhibition of proline specific enzymes by amino acid thioxopyrrolidides and thiazolidides". | | | |
| CE | E Martindale The Extra Pharmacopoeia, 30 th Edition, London Pharmaceutical Press, 1993, Page 1619. | | | |
| CF | AMASHEH, S., et al., "Electrophysiological analysis of the function of the mammalian renal peptide transporter expressed in Xenopus Laevis oocytes". <u>J. Physiol.</u> 504, 169-174 (1997) | | | |
| CG | Arai ET AL., "Synthesis of prolyl endopeptidase inhibitors and evaluation of their structure-activity relationship vitro inhibition of prolyl endopeptidase from Canine Brain" CHEMICAL AND PHARMACEUTICAL BULLE 41, No. 9, 1993, pages. 1583-1588. | | | |
| СН | DURINX, C.; et al.; "Reference Values for Laboratory Parameters". Clin Chem Lab M | | | |
| CI | GOSSRAU, R.; "Cytochemistry of Membra | ane Proteases". Histochem J, 1985, July; | 17 (7):737-71, 1 page. | |
| Cl | HAHN, T.; et al.; "Enzyme Histochemical Cultured Villous Explants from Human Fire page. | | | |
| CK | HEYMANN, E. et al., "Has Dipeptidyl Pep 1984, January, 2;62 (1) :2-10, 1 page. | otidase IV an Effect on Blood Pressure an | d Coagulation." Klin Wochensc | |
| CL | CL Lin, J. et al.: "Inhibition of depeptidyl peptidase IV by fluoroolefin-containing n-peptidyl-O-hydroxylamine peptidomimetics" PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF USA, Vol. 95, November 1998, pages 14020-14024. | | | |
| СМ | KOROM, S., et al "Inhibition of CD26/diperecipients", <u>Transplantation</u> , Vol. 63, 1495 | | ongs cardiac allograft survival in | |
| CN | MAGYAR, C.E. et al., "Proximal Tubule N Hypertension." <u>Am J. Physiol Renal Physi</u> | Na Transporter Responses are the same dod, 2000, August; 279 (2):F358-69, 1 pages | uring Acute and Chronic ge. | |
| СО | Martindale The Extra Pharmacopoeia, 30th | Edition, London Pharmaceutical Press, 1 | 993, Page 36. | |
| iner: | | Date: | | |

| NFOIENA | FORM PTO-1449 TION DISCLOSURE STATEMENT | ATTY. DOCKET NO. 20488/57 | SERIAL NO. 10/667,200 | |
|---------|---|--|--|--|
| | | APPLICANT(S): Kühn-Wache et a | al. | |
| | | FILING DATE: September 18, 2003 | ART UNIT: Not Yet Assigned | |
| | OTHER DOCUMENTS (INCLUDING A | AUTHOR, TITLE, DATE, PERTINE | NT PAGES, ETC.) | |
| СР | MENTLEIN, R., et al., "Proteolytic process Pept. 49, 133 -144 (1993) | ing of neuropeptide Y and peptide YY b | y dipeptidyl peptidase IV". Regul. | |
| CQ | PAPIES, B. et al., "Isoenzyme (Lactate Deh Activity Changes in Blood Plasma Likely In 1991; 33 (3):218-26, 1 page. | | | |
| CR | QURESHI. N.U.; et al., "Endogenous Neuro Shock". Regul Pept, 1998, September 25; | | uring Endotoxic and Hemorrhagic | |
| CS | TANKA, S., et al., "Suppression of arthritis 19, No. 1 Pages 15-24, (1997) | by the inhibitors of dipeptidyl peptidase | e IV". <u>Int. J. Immunopharmacol</u> , Vo | |
| СТ | The Merck Index, 11th Edition, An Encyclo | pedia of Chemicals, Drugs, and Biologic | als, 1989, Page 934 | |
| CU | The Merck Index, 12 th Edition, An Encyclopedia of Chemicals, Drugs, and Biologicals, 1996, Page 1014. | | | |
| CV | Deacon et al., <u>Journal of Clinical Endocrinology and Metabolism</u> , "Degradation of Glucagon-Like Peptide-1 by Hum Plasma in Vitro Yields and N-Terminally Truncated Peptide That Is a Major Endogenous Metabolite in Vivo", (1995 80(3): 952-957. | | | |
| CW | G.G. Duncan, Diseases of Metabolism (Asia | an edition), "Diabetes Mellitus", (1966), | p 951-957. | |
| CX | Gutniak et al., <u>Diabetes Care</u> , "Subcutaneou Postprandial Glycemia in NIDDM", Septen | | cagon-Like Peptide 1 Abolishes | |
| CY | Gutniak et al., New England Journal of Me Normal Subjects and Patients With Diabete | | eagon-like peptide-1 (7-36) Amide in | |
| CZ | H.A. Smith et al., <u>Veterinary Pathology</u> (for (1972), p 1018-1020. | urth edition), "Diseases and Disorders of | Metabolism: Deficiency Diseases", | |
| CAA | Hendrick et al., <u>Metabolism – Clinical and I</u> Rats", January 1993, 42(1): 1-6. | Experimental, "Glucagon-like Peptide-I- | (7-37) Suppresses Hyperglycemia in | |
| CAB | Hoffmann et al., <u>Journal of Chromatograph</u> and non-substrate X-X-Pro- oligopeptides a | | | |
| CAC | Index Nominum, International Drug Directo | ory 1992/1993, Medpharm Scientific Pul | blishers, pages 728-729. | |
| CAD | Mannucci et al., <u>Diabetes Care</u> , "Effect of Metformin on Glucagon-Like Peptide 1 (GLP-1) and Leptin Levels in C Nondiabetic Subjects", 24(3): 489-494, March 2001. | | | |
| CAE | Nauck et al., <u>Diabetologia</u> , "Normalization in Type 2 (non-insulin-dependent) diabetic | | glucagon-like peptide 1 (7-36 amid | |
| CAF | Pauly et al., Regulatory Peptides, "Abstract Peptides", July 15, 1996, 64(1-3): 148 plus | s Issue: Abstracts from the 11 th Internaticover. | onal Symposium on Regulatory | |
| iner: | | Date: | | |

| NFORMA | FORM PTO-1449 TION DISCLOSURE STATEMENT | ATTY. DOCKET NO. 20488/57 | SERIAL NO. 10/667,200 | |
|--------|---|---|---|--|
| | • | APPLICANT(S): Kühn-Wache et al. | | |
| | | FILING DATE: September 18, 2003 | ART UNIT: Not Yet Assigned | |
| | OTHER DOCUMENTS (INCLUDING AU | THOR, TITLE, DATE, PERTINEN | T PAGES, ETC.) | |
| CAG | Stryer, Biochemistry 3 rd Ed., "Protein Conformation of the Biochemistry 3 rd Ed.," | mation, Dynamics, and Function", 1988 | 3, p 191-193. | |
| САН | T.J. Kieffer et al., "Degradation of Glucose-D In Vitro and In Vivo by DP IV", Endocrinology | | Truncated Glucagon-Like Peptide | |
| CAI | The Merck Index, An Encyclopedia of Chemi | cals and Drugs, 9th Edition, Merck & C | Co., Inc., 1976, page 773 | |
| CAJ | C.B. Welch, Medical Management of Non-Instance Association, "Diagnosis and Classification" p | sulin-Dependent (Type II) Diabetes, 3 rd o. 3, 1994, Pharmacologic Intervention | edition, American Diabetes (2 pages). | |
| CAK | WETZEL, W., et al., "Effects of the CLIP frag Neuropeptides, 31, 41- 45 (1997) | gment ACTH 20-24 on the duration of | REM sleep episodes". | |
| CAL | Willms et al., <u>Journal of Clinical Endocrinology</u> Secretion after a Liquid Test Meal: Effects of (Noninsulin-Dependent) Diabetic Patients", 1 | f Exogenous Glucagon-Like Peptide-1 | | |
| CAM | M Ashworth et al., <u>BIOORG. MED. CHEM. LETT.</u> , "2-Cyanopyrrolidides as Potent, Stable Inhibitors of Dipepting Peptidase IV", (1996), 6(10): 1163-1166. | | | |
| CAN | N Badia-Elder N.E. et al., <u>Alcoholism Clinical and Experimental Research</u> , "Effects of Neuropeptide Y (NPY) Intake and Anxiety in High and Low Alcohol Drinking (HAD1/LAD1) Rats", (2000), 24(5): 82A. | | | |
| CAO | C.F. Deacon et al., <u>Diabetes</u> , "Both Subcutane Rapidly Degraded from the NH ₂ -Terminus in 1126-1131. | eously and Intravenously Administered Type II Diabetic Patients and in Health | Glucagon-Like Peptide I Are ny Subjects", September 1995, 44: | |
| CAP | Edwards, J.V. et al., <u>J. Peptide Res.</u> , "Synthes Sequences on Cotton," (1999), 54: 536-543. | "Dipeptidyl peptidase IV (DP IV) and Superoxide Dismutase Activity in Inhibitory Pepdides and Zn ²⁺ in Vitro", (1990), 75(1): 35-44. | | |
| CAQ | Endroczi et al., <u>ACTA PHYSIOL. HUNG.</u> , "I Thymus-Derived Lymphocytes: Effects of Inh | | | |
| CAR | Frohman et al., <u>Journal of Clin. Invest.</u> , "Rapi Plasma in Vitro and in Vivo to a Biologically p 906-913 | id Enzymatic Degradation of Growth H Inactive Product Cleaved at the NH ₂ T | ormone-releasing Hormone by erminus", Volume 78, October 1986 | |
| CAS | Goodman & Gilman's The Pharmacological I | Basis of Therapeutics, Ninth Edition, (1996), p. 1510. ary Peptides Derived from β-Casein," Peptides 21 (2000) 807-809. | | |
| CAT | Hyun Sook Lee et al., "Cathepsin B Inhibitory | | | |
| CAU | Nathan et al., <u>Diabetes Care</u> , "Insulinotropic A Subjects", February 1992, 15(2): 270-275. | Action of Glucagonlike Peptide-1-(7-37 | 7) in Diabetic and Nondiabetic | |
| CAV | Pauly et al., Metabolism, "Improved Glucose Inhibitor Ile-Thiazolidide", (1999), 48(3): 385 | | peptidyl Peptidase IV (CD26) | |
| CAW | Pschyrembel, Kininisches Wörterbuch 257, A | Auflage, (1994), 9 pages. | | |
| ner: | | Date: | | |

| INFOR | FORM PTO-1449 MATION DISCLOSURE STATEMENT | ATTY. DOCKET NO. 20488/57 | SERIAL NO. 10/667,200 | |
|-----------|---|--|-------------------------------------|--|
| ·,· | | APPLICANT(S): Kühn-Wache et al. | | |
| | | FILING DATE: September 18, 2003 | ART UNIT: Not Yet Assigned | |
| | OTHER DOCUMENTS (INCLUDING A | UTHOR, TITLE, DATE, PERTINEN | NT PAGES, ETC.) | |
| CA | Snow et al., <u>Advances In Medicinal Chemistr</u> Immunosuppressive Agents", Vol. 3, (1995), | | dyl Peptidase IV: A New Class of | |
| CA | Thorens et al., <u>Diabetes</u> , "Glucagon-Like Per NIDDM", (1993), 42:1219-1225. | petide-I and the Control of Insulin Secre | etion in the Normal State and in | |
| CA | Z Vidal, (1993), 69 th Edition, p. 612-613. | | | |
| СВА | Wakselman et al., "Inhibition of HIV-1 infec DPP IV activity of CD26", Abstract P 44 of t | | | |
| СВІ | Wettstein, J.G. et al. <u>Pharmacology & Therap</u> (1995), 65(3): 397-414. | peutics, "Central Nervous System Phari | nacology of Neuropeptide Y.", | |
| СВ | Bergmeier, Stephen C., <u>Tetrahedron, Elsevier</u> (2000), pages 2561-2576. | r Science Ltd., "The Synthesis of Vicin | al Amino Alcohols", Vol. 56, No. 17 | |
| СВІ | | lsevier Science Ltd., "Enantioselective Synthesis of $β$ -Hydroxy Amines and rogenation of $α$ -Amido Ketones", Vol. 11, No. 16 (2000), pages 3257-3261. | | |
| CBI | Munglani R. et al., Drugs, <u>Adis International</u> Anxiolytic and Antihypertensive", (1996) 52 | Ltd, At, "The Therapeutic Potential of Neuropeptide Y Analgesic, 2(3): 371-389. | | |
| CB | Orskov, Cathrine et al., "Proglucagon Producin the Fasting State and after Oral Glucose ar | | | |
| СВ | Reinhold, D. et al., <u>Journal of Neuroimmuno</u> Human MBP-Specific CD4 + T Cell Clones | | se IV/CD26 Suppress Activation of | |
| СВІ | H Sengupta, et al., <u>Tetrahedron Letters, Elsevie</u> α-Amino Acylation Reactions: A New Synth 4107-4110. | | | |
| СВ | Stöckel-Maschek, A., et al., <u>Biochimica et Bi</u> Inhibitors of Proline Specific Peptidases", (2 | | yrrolidides and Thiazolidides: new | |
| СВ | Stryer, Lubert, Biochemistry, "Amino Acid I | Degradation and the Urea Cycle" (1975 |) pages 451-452. | |
| CBI | Mentlein et al., <u>Eur. J. Biochem</u> , Dipeptidyl-Peptide-1(7-36)Amide, Peptide Histidine Me 214, pages 829-835. | Peptidase IV Hydrolyses Gastric Inhibitory Polypeptide, Glucagon-Like ethionine and is Responsible for Their Degradation in Human Serum. (1993) | | |
| СВ | Augustyns et al., <u>Eur. J. Med. Chem.</u> , "Pyrro Dipeptidyl Peptidase IV", (1997), vol. 32, pa | | y Relationship as Inhibitors of | |
| СВ | Wen-Tien Chen et al. "Seprase Complexes in (2003). | n Cellular Invasiveness", <u>Cancer and M</u> | letastasis Reviews 22: 259-269, | |
| | | | | |
| Examiner: | | Date: | | |

| INFORMA | FORM PTO-1449 ATION DISCLOSURE STATEMENT | ATTY. DOCKET NO. 20488/57 | SERIAL NO. 10/667,200 | |
|-----------|--|--|---|--|
| | | APPLICANT(S): Kühn-Wache et al. | | |
| | | FILING DATE: September 18, 2003 | ART UNIT: Not Yet Assigned | |
| | OTHER DOCUMENTS (INCLUDING AU | THOR, TITLE, DATE, PERTINEN | T PAGES, ETC.) | |
| CBN | Victor A. Gault et al., "Glucose-Dependent In Treatment of Obesity-Diabetes", <i>Biochemical</i> | | | |
| СВО | Lader, Malcolm H., MD, "Assessment Method <u>Psychopharmacology</u> , (1981), Vol. 1, No. 6, p | | ety", <u>Journal of Clinical</u> | |
| СВР | Winslow, R., "Novartis Drug Alters Picture for | or Diabetes" <u>Wall Street Journal</u> , Wed. | , December 27, 2000, p. B2. | |
| CBQ | Ansorge, S., et al., "Membrane-bound peptida (1991) 4-6, pp. 799-807. | ses of lymphocytes: Functional implica | ations", Biomed. Biochim, Acta 50 | |
| CBR | Dodge, R.W., et al., "Folding and Unfolding I Ribonuclease A," <i>Biochemistry</i> 1996, 35, pp. | | ants of Bovine Pancreatic | |
| CBS | Hans-Ulrich Demuth, "Recent Developments Vol. 3, pp. 249-278. | in Inhibiting Cysteine and Serine Prote | ases", <u>J. Enzyme Inhibition</u> , 1990, | |
| СВТ | BT Gomez, S., et al., "Relationship between endo- and exopeptidases in a processing enzyme system: Activation of a endoprotease by the aminopeptidase B-like activity in somatostatin-28 convertase", <i>Proc. Natl. Acad. Sci. USA</i> , \pp. 5468-5472, August 1988. | | | |
| CBU | Hegen, M., et al., "The T Cell Triggering Mol <u>The Journal of Immunology</u> , vol. 144, pp. 290 | e amyloid A4-generating enzyme as a prolyl endopeptidase," Federation of | | |
| CBV | Ishiura, S., et al., "Identification of a putative European Biochemical Societies, Vol. 260, No. | | | |
| CBW | Kräusslich, Hans-Georg, et al., "Viral Proteina | ases", <i>Ann. Rev. Biochem</i> . 1988, 57 pp. | 701-754. | |
| CBX | Pederson, R.A., et al., "Improved Glucose Tol Peptidase IV Inhibitor Isoleucine Thiazolidide | | | |
| СВҮ | Vanhoof, G., et al., "Proline motifs in peptide pp. 736-744. | s and their biological processing", The | FASEB Journal, Vol. 9, June 1995, | |
| CBZ | Walter, R., et al., "Proline Specific Endo- and 18, 1980, pp. 111-127 | Exopeptidases", Molecular & Cellular | <i>Biochemistry</i> , Vol. 30, No. 2, April | |
| СВА | Kessler, Von Horst, "Konformation und biolo 509-520. | logische Wirkung von cyclischen Peptiden", Angew. Chem. 94 (1982) pp. | | |
| ССВ | Kirschke, H. et al., "Proteinases 1: Lysosomal 1634. | Cysteine Proteinases" <u>Protein Profile</u> , | Vol. 2, Issue 14, 1995, pp. 1583- | |
| ccc | Yaron, A., et al., "Proline-Dependent Structur <u>Biochemistry and Molecular Biology</u> , 28(1), p | | s and Proteins" <u>Critical Reviews in</u> | |
| CCD | Vallee et al., "Larval Development of Tribolic Database CAPLUS on STN, Accession No.: 1 | | | |
| Examiner: | | Date: | | |